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(54) Title: **BIOMEDICAL ELECTROCHEMICAL SENSOR ARRAY AND METHOD OF FABRICATION**

(57) Abstract: Methods for fabricating a plurality of sensors on a flexible substrate, with each sensor having associated electrodes and at least one well include (a) providing a flexible substrate material layer having a surface area defined by a length and width thereof; (b) forming a plurality of sensor elements onto the flexible substrate material layer, each sensor element comprising at least one metallic electrode; (c) disposing at least one coverlay sheet layer over the flexible substrate sandwiching the sensor elements therebetween; (d) laminating the at least one coverlay sheet layer having an associated thickness to the flexible substrate; and (e) removing predetermined regions of the laminated coverlay sheet layer from the flexible substrate layer to expose a portion of the underlying metallic pattern of each sensor element and to define a well with a depth corresponding to the thickness of the coverlay sheet layer. The disclosure also describes multi-layer laminated flexible sensors and arrays of sensors with wells having enhanced well capacity and/or depth.

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BIOMEDICAL ELECTROCHEMICAL SENSOR ARRAY AND METHOD OF FABRICATION

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Work related to the invention was sponsored by the NSF CECT under Grant No. CDR-8622201. The United States Government has certain rights to this  
5 invention.

Related Applications

This application claims priority to U.S. Provisional Application Serial Number 60/332,194, filed November 16, 2001, the contents of which are hereby incorporated by reference as if recited in full herein.

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Field of the Invention

The present invention relates to methods for fabricating electrochemical sensors or wells on flexible substrates and associated products. The products may be  
15 particularly suitable for use as disposable biomedical sensors.

Background of the Invention

Probes or sensors used in medical diagnostic or evaluation procedures often use electrochemical detection provided by dry or fluid/liquid chemistries/electrolytes  
20 placed on top of electrodes formed of precious metals (gold, platinum, etc). The probes or sensors can employ chemistries/electrolytes such as solid potassium chloride (such as for reference electrodes) or other chemicals, hydrogels (sometimes containing an internal electrolyte underneath the membrane of an ion-sensitive electrode), or enzyme-containing material. The sensors or probes also typically  
25 employ wells or small pools, some of which can be configured to act as capillary spaces to guide quantities of a sample solution (such as blood) to and/or from the electrodes on the probe or sensor.

For many of these applications, the wells are patterned into materials which are selected so that they are compatible with flexible substrates such as polyimide films (Kapton®, Upilex®, and the like). In the past, thin film processing techniques have had problems generating coatings thick enough for proper well formation in chemical sensor applications. In addition, screen printed materials used with thick film processing techniques may be either incompatible with flexible materials or inhibit the formation of fine line resolution desired for small or miniaturized electrodes.

Cosofret et al., in *Microfabricated Sensor Arrays Sensitive to pH and K<sup>+</sup> for Ionic Distribution Measurements in the Beating Heart*, 67 Anal. Chem., pp. 1647-1653 (1995), described spin-coating a polyimide layer of about 30µm onto a film or substrate. Unfortunately, spin-coating methods can, as a practical matter, limit the well depth and/or precise boundary or perimeter definition during formation. In addition, spin-coating methods may be limited to batch fabrication processes and are generally not commercially compatible with high volume, low-cost (continuous or semi-continuous) mass production methods. In view of the foregoing, there is a need for improved, economic ways to fabricate wells and microenvironments for electrochemical sensors on flexible substrates.

#### Summary of the Invention

In certain embodiments, the present invention is directed to methods for fabricating a plurality of sensors on a flexible substrate, each sensor having at least one associated electrode and at least one well. As used herein, the term “well” means a reservoir or chamber used to receive or hold a quantity of fluid therein (typically sized and configured as a microfluidic environment). As such, the term “well” includes at least one discrete chamber or a plurality of chambers (in fluid communication or in fluid isolation, as the application desires) and can alternatively or additionally include one or more channels (linear or other desired complex or irregular shapes (such as spiral, annular, etc.)), or combinations of a well(s) and channel(s).

In certain embodiments, the method includes: (a) providing a flexible substrate material layer having a surface area defined by a length and width thereof; (b) forming a plurality of sensors onto the flexible substrate material layer, each sensor

comprising a predetermined metallic pattern defining at least one electrode; (c) disposing at least one coverlay sheet over the flexible substrate sandwiching the sensors therebetween, the coverlay sheet having an associated thickness; (d) laminating the at least one coverlay sheet to the flexible substrate layer; and (e)  
5 removing predetermined regions of the laminated coverlay sheet from the flexible substrate layer to define a well (which may be or include a channel) with a depth corresponding to the thickness of the coverlay sheet.

In certain embodiments, the removing step also exposes a portion of the underlying metallic pattern of each sensor (such as bond pads and an interdigitated  
10 array or "IDA"). The array of sensors can be arranged such that the sensors are aligned back to back and side by side to occupy a major portion of the surface area of the flexible substrate. In addition, the patterned coverlay can be configured such that the well is a microfluidic channel or a channel with a well. In certain embodiments, the assembly may be configured such that there are openings in the coverlay for bond  
15 pads and the like to make any desired electrical connection(s).

Other embodiments of the invention are directed to arrays of flexible sensors. The arrays of flexible sensors include: (a) a flexible substrate layer having opposing primary surfaces, (b) an electrode layer disposed as a repetition of metallic electrically conductive patterns on one of the primary surfaces of the substrate layer, the metallic  
20 pattern corresponding to a desired electrode arrangement for a respective sensor; and (c) a first coverlay sheet layer having a thickness overlying and laminated to the first flexible substrate layer to sandwich the electrode layer therebetween. The third coverlay sheet layer has a plurality of apertures formed therein. The apertures define a well for each of the sensors on the flexible substrate. The wells have a depth  
25 corresponding to the thickness of the coverlay sheet layer.

Other embodiments are directed to flexible sensors, which can be single use or disposable bioactive sensors. Similar to the array of sensors, the individual sensors can be multi-layer laminated structures including: (a) a flexible substrate layer; (b) an electrode layer comprising a conductive pattern of material disposed onto one of the  
30 primary surfaces of the first flexible substrate layer; and (c) a first flexible coverlay layer overlying the electrode layer and laminated to the electrode layer and the substrate layer, wherein the first flexible coverlay layer has a well formed therein, the well having a depth of at least about 1-10 mils (.001-.01 inches) or, in a metric

system, at least about 25-250  $\mu\text{m}$ . Of course greater well depths can also be generated, such as by using thicker coverlay sheets or combinations of sheets, to yield well depths of about 12 mils (about 300  $\mu\text{m}$ ) or more, depending on the application.

5 In certain embodiments, the array of sensors or each sensor can include a second coverlay layer having a thickness of between about 1-10 mils overlying and secured to the first coverlay layer. The second coverlay layer also has a plurality of apertures formed therein, the apertures corresponding to the apertures in the first coverlay layer. Thus, the wells have a depth corresponding to the combined thickness of the first and second coverlay layers. In other embodiments, a third coverlay layer  
10 can also be employed by laminating it to the second coverlay layer and removing the material overlying the well site to provide a well depth corresponding to the thickness of the first, second, and third coverlay layers.

The method of fabricating the sensor arrays can be carried out in an automated continuous production run that increases the production capacity over batch type  
15 processes. In addition, the wells can be formed with increased volume, capacity, or depth over conventional microfabrication techniques. The method can be performed such that the sensors are arranged on the flexible substrate in a high-density pattern of at least about 4 sensors per square inch when measured over about 122 square inches. In other high-density embodiments, for a sheet which is 12 inches by 12 inches (144  
20 square inches), about 750 sensors can be arranged thereon, averaging at least about 5 sensors per square inch. In certain embodiments, the sensors and arrays are configured to be heat resistant or to withstand sterilization procedures suitable for biomedical products.

The coverlay material can be a photosensitive film such as a dry film material.  
25 Examples of suitable coverlay materials include photoimageable polymers, acrylics, and derivatives thereof including, but not limited to, commercially available PYRALUX® PC and VACREL® from DuPont, and CONFORMASK® from Morton. In addition, the coverlay sheet may be a pre-laminated sheet of a plurality of plies of one or more types and/or varying thickness of dry film coverlay materials and  
30 may also include desired coatings.

The foregoing and other objects and aspects of the present invention are explained in detail in the specification set forth below.

### Brief Description of the Drawings

**Figure 1** is a flow chart of method steps for fabricating sensors with wells on a flexible substrate according to embodiments of the present invention.

**Figures 2a-2g** are side views of a fabrication sequence of flexible substrate sensors using coverlay sheet material according to embodiments of the present invention.

**Figures 3a-3g** are top views of the sequence shown in **Figures 2a-2g**, (with **Figures 2a** and **3a** correspond to one another, **Figures 2b** and **3b** corresponding to one another and so on.

**Figure 4** is a flow chart of the sequence of fabrication steps illustrated in **Figures 2** and **3**.

**Figure 5** is a photocopy of the upper surface of a partial sheet of an array of sensors with wells on a flexible substrate according to embodiments of the present invention.

**Figure 6a** is a greatly enlarged side perspective view of a sensor with a well according to embodiments of the present invention.

**Figure 6b** is a greatly enlarged side perspective view of a sensor with a well having a depth corresponding to the combined thickness of multiple coverlay sheets.

**Figure 7** is a top view of a partial sheet of an array of sensors drawn to scale according to embodiments of the present invention.

**Figures 8a-8i** are schematic illustrations of stations in a production line for fabricating arrays of flexible sensors according to embodiments of the present invention.

### Description of Embodiments of the Invention

The present invention will now be described more fully hereinafter with reference to the accompanying figures, in which preferred embodiments of the invention are shown. This invention may, however, be embodied in many different forms and should not be construed as limited to the embodiments set forth herein. Like numbers refer to like elements throughout. In the figures, layers, components, or features may be exaggerated for clarity.

In certain embodiments, as shown in **Figure 1**, steps for fabricating a flexible sensor can include first providing a flexible substrate material layer (**Block 100**). The

flexible substrate material layer can be an elastomeric film such as polyimide films. Examples of commercially available films include, but are not limited to, Kapton®, Upilex®, Kaladex® and the like. Next, a plurality of sensors can be formed on the flexible substrate layer via depositing or forming a desired conductive and/or metallic electrode pattern thereon (**Block 110**). Any suitable metal trace fabrication technique may be employed, such as sputtering or deposition of the metal followed by photolithography or laser ablation, stenciling, screen-printing, shadow masking, and the like.

In any event, after the metallic pattern is formed on the substrate, at least one coverlay sheet layer having an associated thickness can then be disposed to overlie the flexible substrate layer so as to sandwich the sensors or metallic pattern therebetween (**Block 120**). The coverlay sheet can be a photosensitive and/or photoimageable coverlay dry film material. Examples of suitable coverlay materials include photoimageable polymers, acrylics, flexible composites, and derivatives thereof including, but not limited to, commercially available PYRALUX® PC and VACREL® from DuPont, and CONFORMASK® from Morton. In addition, the coverlay sheet may be a pre-laminated sheet of a plurality of plies of one or more types and/or varying thickness of dry film coverlay materials and may also include desired coatings. The coverlay material may be selected so as to be heat resistant or compatible with irradiation sterilization procedures as, in use, the sensor may be exposed to sterilization procedures, particularly for biomedical applications.

The coverlay sheet is laminated to the underlying flexible substrate layer (*i.e.*, the layers are united) (**Block 130**). The layers can be united by hot roll lamination techniques, or other suitable lamination means suitable to unite the layers together. Predetermined regions of the laminated coverlay sheet can be removed from the flexible substrate layer to define a well with a depth corresponding to the thickness of the coverlay sheet layer (**Block 140**). In certain embodiments, two or more coverlay sheets can be laminated, serially, onto the flexible substrate to define a well with a depth corresponding to the combined thickness of the coverlay sheets used.

Figures 2 and 3 illustrate a sequence of operations, suitable for certain embodiments, which can be used to form a flexible sensor array (which can be separated to form individual disposable sensors as will be discussed further below). As shown in Figure 2a and 3a, first a conductive metal 20 can be deposited onto the

flexible substrate layer 10 (**Block 150, Figure 4**). The conductive metal 20 can be any suitable conductor as is well known to those of skill in the art, including, but not limited to, gold, silver, platinum, palladium, titanium, chromium and mixtures thereof. In certain embodiments, the metal 20 can be formed as a relatively thin layer of about 30-200nm, and is typically about 100nm. The metal 20 can be applied via a sputtering process. A suitable metal coated flexible substrate material is available from Techni-Met, Inc., located in Windsor, CT. Additional additives such as adhesive enhancing materials (such as seed layers of chromium or titanium) can be sputtered or sprayed or otherwise deposited onto the substrate 10 to promote the adhesion of the metal 20 to the substrate layer 10.

As shown in **Figures 2b and 3b**, a dry film photoresist material layer 30 can be laminated onto the metal layer 20 (**Figure 4, Block 160**). The photoresist material 30 can be a dry film resist such as Riston® CM206 from DuPont Industries. The photoresist material 30 may be wet laminated by hot rolling the photoresist onto the metal coated surface 20 of the substrate using a HRL-24 hot roll laminator (also available from DuPont). Subsequently, as shown in **Figure 2c**, a photomask 35 with a predetermined mask pattern is positioned between the photoresist material 30 and an ultraviolet light source and the ultraviolet light 38 travels through desired apertures in the mask 35 to expose photoresist material about the unmasked regions 39e but not the masked regions 39u, thereby forming a desired exposure pattern in the laminated dry film photoresist material 30. Thus, predetermined regions of the photoresist material are exposed (**Figure 4, Block 170**). In the embodiment shown, a negative photoprocessing system is employed, the mask being configured to cover the regions (and prevent exposure) which will define the desired trace or electrode pattern. However, a positive imaging or photoprocessing technique can also be used as is well known to those of skill in the art.

As shown in **Figures 2d and 3d**, the photoresist material 30 is then developed (**Figure 4, Block 180**), leaving the patterned photoresist 39p on the surface (the exposed regions being removed). **Figures 2e and 3e** illustrate that next the metal pattern is etched into the metal layer 20 corresponding to the laminated photoresist material remaining thereon (**Figure 4, Block 190**). As shown in **Figures 2e and 3e**, after the metal pattern 20p is formed, the remainder of the photoresist material 30 is



removed or stripped (**Figure 4, Block 195**), leaving the substrate layer **20** with the desired metallic pattern **20p** thereon.

Next, as shown in **Figures 2f** and **3f** (and **Figure 4, Blocks 200, 205**), a coverlay material **40** is laminated onto the flexible substrate **10** sandwiching the metallic pattern **20p** therebetween. The lamination can be performed as a hot roll process, which presses the two layers together. The lamination can be carried out under vacuum to help remove air which may be trapped or residing between the two joining surfaces.

As before, **Figures 2f** and **3f** illustrate that photolithography can be used to expose desired regions **40e** of the coverlay material **40**, which is then selectively removed (or if desired, the reverse processing can be used, *i.e.*, the unexposed material can be removed) thereby forming the desired pattern in the coverlay material **40p**, which then forms the well and/or exposes the underlying metal **20** or substrate material **10**. The exposed material is removed about a well region formed by the selective removal of the coverlay material **40**, the well **40w** having a depth corresponding to the thickness of the coverlay. As shown in **Figure 2f**, a mask **45** is positioned between an ultraviolet light source and the light rays travel through openings in the mask **45** to expose the predetermined regions in the coverlay material **40**. The coverlay material is then developed (**Figure 4, Block 210**) and the unwanted coverlay material **40** is removed or stripped, leaving a flexible array of sensors with surface regions exposed to the underlying material (either the flexible substrate **10**, or the metal layer **20**). The coverlay material can then be cured (**Figure 4, Block 215**). Any suitable curing technique can be employed, but typically a thermal curing process is employed to heat the coverlay material to desired temperatures and for desired cure times to thermally cross-link the coverlay material. Doing so “permanently secures” the coverlay material(s) to the underlying substrate and/or to inhibit the degradation or separation of the structure in use (maintaining the integrity of the attachment between the coverlay material and the underlying materials). Other curing means may be used, as suitable, depending on the materials employed, including, but not limited to, oven or other heat source, microwave, RF, or ultrasound energy, and/or laser, ultraviolet light or other light source.

**Figure 5** illustrates an example of an array of flexible sensors **50** according to certain embodiments of the invention. As shown, the array **50** can be fabricated so as

to provide a plurality of sensors 55. In certain embodiments, adjacent sensors 55 are arranged side-by-side and back-to-back or otherwise oriented to provide a high density arrangement of sensors and/or to occupy a major portion, and typically substantially all, of the surface area of the flexible array 50. The term "high density" means that, when measured over at least about  $121\text{in}^2$  (typically in a square sheet of about  $11 \times 11$  inches), on average, the array 50 holds about 490 sensors 55 or at least about 4 complete sensors 55 per square inch. In certain embodiments, when measured over a  $12 \times 12$  inch area (or a  $144\text{in}^2$ ), the array can hold about 750 sensors, or at least about 5 sensors on average. Figure 7 illustrates an exemplary layout of a partial sheet of an array 50' of sensors drawn to scale; the array 50' can be produced with at least 35 sensors in the vertical direction and 14 sensors in the horizontal direction (the  $35 \times 14$  array shown corresponds to an  $11 \times 11$  sheet). Of course, other array sizes can be used and the sensors themselves further miniaturized or enlarged, depending on the desired application.

As shown in Figure 5, the sensors 55 can be symmetrically arranged in columns (the column width corresponding to the length of the sensor 55l) and rows (with the row height corresponding to the width of the sensor 55w), with gap spaces positioned adjacent each sensor 55 to facilitate separation from the array 50 for individual use so as to inhibit damage to the adjacent sensor upon removal. The array 50 can include alignment marks 59 that can facilitate the alignment of masks and system components with the metal pattern of individual sensors or groups of sensors on the array 50 during fabrication.

It is noted that the sensors 55 on the array 50 (including the depth of the well 40w and metal pattern 20p which defines the desired electrical or electrode arrangement) can be alternately configured, shaped, and arranged. For example, the sensor length 55l can be disposed vertically on the flexible substrate 10 such that the row height corresponds to the length of the sensor or the electrode can include curvilinear traces or circular, triangulated, or other electrode shapes. In Figure 5 and 7, the darkest regions correspond to locations where the laminated overlay was removed from the underlying flexible substrate, exposing either the underlying substrate or a portion of the metallic pattern 20p. The lighter regions correspond to locations where the laminated coverlay material 40 remains intact (sandwiching a portion of the underlying metal pattern 20p to the flexible substrate 10).

As shown in **Figure 5**, the array **50** is planar and includes a metallic pattern formed on one primary surface of the underlying flexible substrate **10**, each sensor **55** including a metallic pattern which defines the arrangement of at least one electrode **60e**. In the embodiment shown in **Figures 5-7**, the electrode **60e** includes two spaced apart bond pads **60p**, each having a respective connecting trace **60t**, and an interdigitated array (IDA) **60i**. In certain embodiments, the IDA can have a structure width which may be in the sub- $\mu\text{m}$  range. In the embodiment shown, the IDA **60i** is positioned in the well **40w**. For additional information on IDA's, see, e.g., U.S. Patent No. 5,670,031 and WO 97/34140, the contents of which are hereby incorporated by reference as if recited in full herein. See also, Niwa et al, *Fabrication and characteristics of vertically separated interdigitated array electrodes*, J. Electroanal. Chem., 267, pp. 291-297 (1989); Koichi Aoki, *Theory of the steady-state current of a redox couple at interdigitated array electrodes of which pairs are insulated electrically by steps*, J. Electroanal. Chem. 270, pp. 35-41 (1989); Koichi Akoi, *Quantitative analysis of reversible diffusion-controlled currents of redox soluble species at interdigitated array electrodes under steady state conditions*, J. Electroanal. Chem. 256, pp. 269-282 (1988); and Horiuchi et al., *Limiting Current Enhancement by Self-Induced Redox Cycling on a Micro-Macro Twin Electrode*, J. Electrochem. Soc., Vol. 138, No. 12 (Dec. 1991). The contents of these documents are also hereby incorporated by reference as if recited in full herein.

**Figure 6a** illustrates a greatly enlarged sensor **55**. As shown, the sensor **55** includes a well **40w** having a depth "D" which corresponds to the thickness of the coverlay sheet(s) **40** laminated to the flexible substrate **40**. **Figure 6b** illustrates that the well **40w** can have a depth "D" corresponding to the combined thickness of two laminated coverlay sheets, **40a**, **40b**. As noted above, one, two, or three or more sheets **40** can be used to generate the desired well depth. The sheets (shown in **Figure 6b** as **40a**, **40b**) can have the same or a different thickness and/or can be formed from the same material(s) or different material(s). For example, three sheets, each having a thickness of about 4 mils, can be used to define a well **40w** with a 12 mil depth. Alternatively, a 3 mil and 2 mil sheet can be used to provide a well **40w** with a 5mil depth.

As also noted above, the coverlay sheet **40** can be a pre-laminated sheet of a plurality of plies of materials with or without coatings. In addition, as also noted

above, the coverlay sheet(s) 40 can have a thickness of at least 0.5-10 mils (about 12-250  $\mu\text{m}$ ) and preferably has a thickness of about 1-20 mils (about 25-500  $\mu\text{m}$ ) or more. In certain embodiments, the wells 40 have a depth which is in the range between about 5-15mils.

5 In certain embodiments, the coverlay sheet(s) 40 are selected to define a well depth "D" and perimeter shape 40s which is consistent from sensor to sensor 55 to provide a consistent testing space or volume. This can allow for improved meting of the biological fluid undergoing analysis, thus helping to provide a more consistent sample size to combine with the electrochemical formulation or solution or chemical  
10 substance(s) which may also be contained in the well 40w (not shown). In turn, reducing variation in the sensor operation can promote more reliable test results. Additional description of electrodes and analyte formulations are found in co-pending U.S. Patent Application identified by Attorney Docket No. RDC0002/US, entitled  
15 "ELECTRODES, METHODS, APPARATUSES COMPRISING MICRO-ELECTRODE ARRAYS", the contents of which are hereby incorporated by reference as if recited in full herein.

As shown in Figure 6a, the well 40w can be in fluid communication with an IDA 60i or electrode 60e of a desired configuration which is in electrical communication with opposing electrical connecting traces 60t. Typically, the well  
20 40w has a configuration which opens the laminated coverlay 40 to expose the underlying IDA or electrochemical active components while defining the perimeter shape 40s in a precise repeatable manner to generate a consistent reliable testing environment part to part.

As shown in Figure 6b, the well can be in fluid communication with a  
25 capillary segment 240c having a depth "D" (which is typically the same depth as the well) which directs fluid from the well 40w through the capillary segment 240c to the active test well 240w. The test well 240w may be configured to house an IDA as noted above, or other desired electrical component or electrode 60e, and/or a chemistry formulation corresponding to the test protocol for the particular sensor  
30 application. The sensor 55 can also include one or more electrical traces 60t and one or more bond pads 60b configured, in operation, to be electrically engageable with a testing device capable of receiving and analyzing the signal of the sensor 55 (not shown).

In certain embodiments, the testing device can be a home unit and the sensor can be a disposable (typically, a single use disposable) sensor suitable for use by a patient, for example, to monitor glucose or other analyte levels (or the presence or absence of substances) in the blood or other body fluid or sample. It will be appreciated that the shape, length and configuration of the electrode or metallic pattern as the well shape, configuration or depth can vary depending on the desired end application.

Turning now to **Figures 8a-8i**, an exemplary embodiment of a production line with nine production workstations is shown. In certain embodiments, the production line can be configured to be automated and semi-continuous or continuous. The word “continuous” means that a production run of a desired quantity or length of material can be processed serially through each of the stations and is operated generally without substantial time delay or disruption between stations (*i.e.*, certain delays are expected such as for set-up, tool change, material introduction, maintenance, queues at equipment, shift changes, planned and unplanned downtime, etc). The term “semi-continuous” as used herein, means that fabrication of the array of sensors is carried out by maintaining the product on reels of desired “continuous” lengths through selected stations. Typically, the reels of material are sufficient in length to have continuous production runs through at least the lamination and patterned coverlay stations (shown as stations 6-8). Of course, the in process or processed reels at any particular station may be queued or stored for the next workstation depending on capacity, orders, and the like. That is, for the embodiment shown, reels of flexible substrates and photoresist material as well as coverlays can be used to automatically run the processing steps in each station to form the patterned coverlay laminated to the flexible substrate and electrode surface, preferably even through any desired final curing station (shown as station 9, **Figure 8i**). In addition, the flexible substrate may be cut to form individual sheets at selected points during the process. However, by using reels of arrays or materials, the fabrication process can be automated and run in continuous lengths of material in relatively long production runs (in contrast to batch mode production operations).

Although shown as nine separate workstations in **Figures 8a-8i**, for ease of discussion, it is noted that some of the stations can be multipurpose or combined with other workstations. For example, one or more of stations 2 and 6, stations 3 and 7,

and/or stations 4 and 8 may be configured to be the same physical station or equipment, but configured to laminate, expose, or remove the appropriate material, depending on the desired processing step for that particular product run. In addition, the material is illustrated as being rerolled onto reels at the end of each workstation; however, alternatively, certain or all of the stations can be arranged to directly feed the material to the next workstation (which can be located downstream of the previous workstation) to provide direct material throughput without rolling onto reels station to station. Also, the gold or metal deposited flexible substrate may be pre-fabricated or obtained from a supplier and patterned locally by the process described at workstations 2-9.

For clarity, the workstations have been identified with feature numbers which correspond to the method steps shown in **Figure 4** (*i.e.*, workstation 150s corresponds to the workstation where the method step 150 of depositing gold onto the flexible substrate can be carried out).

Turning now to **Figure 8a**, as shown, a roll coating station 150s conveys or pulls (or otherwise processes) the flexible substrate material 10 so as to coat a selected primary surface with the desired metal (such as gold). As shown, a reel of flexible substrate 10r (shown as Kapton®) is processed to include the metal (labeled as Au) on one of the primary surfaces. The metal coated substrate material is then rolled into a reel 110r. As shown in **Figure 8b**, a laminating station 160s, takes the material reel 110r and combines it with a photoresist film ("PR") 30 to laminate to the underlying metallized surface of the flexible substrate 10 by pressing the materials together via rollers in a hot roll-laminating machine at selected pressures and temperatures; P; T. The laminated photoresist 30, metal 20, and flexible substrate 10 are then rolled onto reel 120r.

**Figure 8c** illustrates an exposing station 170s, where material reel 120r is unrolled and the PR 30 exposed to the ultraviolet light 38 through a mask 35 with a predetermined exposure pattern and the material is re-rolled onto a reel 120r<sub>e</sub> which now holds the exposed PR, metal, and flexible substrate. The material on reel 120r<sub>e</sub> is then introduced to a developing station 180s, as shown in **Figure 8d** where the PR is developed, rinsed and dried, resulting in a composition of patterned PR on metal and flexible substrate which can be rolled onto a reel 125r. As shown in **Figure 8e**, the material on reel 125r can then be introduced to an etching station 190s where the

metal can be etched, the PR stripped, and the patterned metal 20p on flexible substrate 10 dried and rolled onto reel 126r.

Figure 8f illustrates a coverlay application station 200s, where the coverlay sheet 40 can be introduced as a substantially continuous reel of material 40r and the flexible substrate with the metallic pattern can also be introduced as a continuous reel of material. The continuous lengths of materials can be forced together so that the coverlay sheet 40 is secured to the underlying materials. As shown, the coverlay sheet is laminated or united to the patterned metal surface 20p and the underlying flexible substrate in a hot roll lamination machine which presses the two materials together (which may be evacuated during the procedure to reduce the likelihood that air is trapped between the layers). The laminated coverlay 40, metal pattern 20p, and flexible substrate 10 can then be rolled onto a reel 127r. Although not shown, step 6 can be repeated as desired to laminate additional coverlay material sheets onto the first laminated coverlay surface.

Figure 8g illustrates a coverlay exposure station 205s, where the laminated coverlay material 127r is exposed to a light source (similar to the photoresist material processed at station 3) and collected and rolled to form a reel 127r<sub>e</sub> of exposed coverlay material on patterned metal 20p and flexible substrate 10. The reel exposed coverlay material 127r<sub>e</sub> is then taken to developing station 210s (Figure 8h), where the coverlay material 40 is developed to yield a laminated patterned coverlay layer 40p overlying a patterned metal surface 20p on the flexible substrate 10 which can be collected on a reel 128r. The reel of material 128r can then be directed to travel through a thermal curing station 205s to cure the coverlay material 40 (for example, an oven at about 160°C with a conveying tension and speed set so that the coverlay sheet is cured for about 1 hour). Of course, other temperatures and times (and related conveyor speeds can also be used). In addition, a more complex array continuous travel pattern (not shown) can be used to occupy more of the space in the oven (*i.e.*, spiral or zig-zag to use more vertical space). Alternatively, the array 50 can be cut into desired lengths and conveyed through the oven. As shown, the continuous length of array 50 is rolled onto a finished array material spool or reel 129r which can be transported to a pharmaceutical location where a desired chemical formulation can be added to the wells and the sensors split into individual units.

It is also noted that, in certain embodiments, an additional coverlay layer or layers can be positioned to define a ceiling or lid over the underlying laminated coverlay defining the well(s) (not shown). The ceiling coverlay layer can be configured to enclose the underlying surface or portions of the surface such as to enclose the well. The enclosed well configuration may be particularly suitable for enclosed microfluidic testing environments. The ceiling coverlay layer may be patterned to define a port or openings in the ceiling layer to allow electrical or fluid access to desired regions thereunder. In certain embodiments, a port can be patterned into the ceiling coverlay to allow fluid passage to a portion of the well. In other embodiments using enclosed well (chamber and/or channel) configurations, the fluid travel passage can be provided through vias or passages formed up through the substrate layer or formed laterally through an intermediate layer (such as, when viewed from the top, a lateral passage extending from an open end region to the testing well). In these embodiments, an additional ceiling forming set of stations (similar to those used to form the coverlay(s) defining the wells onto the substrate) can be used to laminate the ceiling coverlay to the underlying structure and/or pattern the ceiling coverlay as desired.

The invention is explained in greater detail in the following non-limiting examples.

#### Examples

The following process was used to prepare an article according to embodiments of the invention. According to the method, a gold film or layer is deposited onto a flexible substrate formed of 7 mil thick Kaladex® film using a planar DC magnetron sputtering process and equipment operated Techni-Met Inc. (a roll coating company), located in Windsor, CT. The thickness of the gold film can range from 30 to 200 nm, with a preferred thickness being about 100nm. Seed layers of chromium or titanium can be sputtered between the substrate film and the gold layers to promote better adhesion of the gold to the substrate film; however, gold layers sputtered directly onto the substrate film without such seeding can exhibit sufficient adhesion. Plasma treatment of substrate surface can improve the adhesion of gold.

After the gold was applied to the flexible substrate, a dry film photopolymer resist was laminated to the gold/substrate film. A dry film resist such as that sold under the trademark Riston® CM206 (duPont) was used. The Riston® CM206



photoresist was first wet laminated onto the gold surface of 12" x 12" gold/substrate panels using a HRL-24 hot roll laminator (from duPont). The sealing temperature and lamination speed were about 105°C and 1 meter per minute, respectively. The laminated panel was placed in a Tamarack model 152R exposure system, from  
5 Tamarack Scientific Co., Inc., Anaheim, CA. The release liner was removed from the top surface of the photoresist. A glass/Cr photomask was produced by Advance Reproductions Corporation, North Andover, MA. The Cr side of the mask was treated with an antistick coating (Premitech Inc., Raleigh, NC), and was placed directly onto the photoresist surface of the panel. The laminated panel was exposed to  
10 ultraviolet light of 365 nm through the photomask using an exposure energy of 60 mJ/cm<sup>2</sup>. Unexposed photoresist was stripped from the panel in a rotary vertical lab processor (VLP-20), Circuit Chemistry Equipment, Golden Valley, MN, using 1% potassium carbonate, at room temperature, for 30 seconds using a nozzle pressure of 34 psi. Exposed gold on the sheet was then stripped using an etch bath containing a  
15 solution of 4 parts I<sub>2</sub>:1 part KI:40 parts water vol./vol.; and 0.04 gram Fluorad™ fluorochemical surfactant FC99, (3M, St. Paul, MN) per 100 gram solution, added to the bath to ensure wetting of the gold. Air was bubbled through the bath during the etch process to obtain a sufficiently uniform agitation of the bath mixture. The panel was rinsed with deionized water and residual Riston® CM206 was removed in a 3%  
20 KOH bath.

Articles were fabricated using dry film photoimageable coverlay materials such as that sold under the trademark Vacrel® 8140 (and related series) from duPont or Pyralux® PC series (duPont). The chamber dimensions can be accurately defined by flex circuit photolithography. Depth of the chamber was controlled by the  
25 thickness of the coverlay materials used and/or whether single or multiple layers of the coverlay dry film were used. Chamber depth was achieved by sequential lamination of different coverlay materials as follows: four mil thick Vacrel® 8130 was first laminated to the electrode side of the substrate using a HRL024 (duPont) heated roll laminator at room temperature, using a roller speed of 1 meter per minute.  
30 The electrode panel was then vacuum laminated in a DVL-24 vacuum laminator (duPont) using settings of 120°F, 30 second vacuum dwell, and a 4 second pressure dwell to remove entrapped air between the coverlay film and the electrode substrate. Two mil thick Vacrel® 8120 was laminated next to the Vacrel® 8130 surface using

the HRL-24 at room temperature, with a roller speed of 1 meter/min. The panel was then vacuum laminated again in the DVL-24 vacuum laminator using a 30 second vacuum dwell, 4 second pressure, to remove entrapped air between the two coverlay films.

5           The laminated electrode sheet was placed in the Tamarack 152R system and was exposed to ultraviolet light at 365 nm through the photomask for 22 seconds using an exposure intensity of 17 mW/cm<sup>2</sup>. The unexposed coverlay was stripped from the panel using the VLP-20 Circuit Chemistry Equipment) in 1% K<sub>2</sub>CO<sub>3</sub>, at 140°F, for 75 seconds using a nozzle pressure of 34 psi. The developed laminate  
10 structure was rinsed in deionized water, and then cured at 160°C for 1 hour to thermally crosslink the coverlay material.

          The foregoing is illustrative of the present invention and is not to be construed as limiting thereof. Although a few exemplary embodiments of this invention have been described, those skilled in the art will readily appreciate that many modifications  
15 are possible in the exemplary embodiments without materially departing from the novel teachings and advantages of this invention. Accordingly, all such modifications are intended to be included within the scope of this invention as defined in the claims. In the claims, means-plus-function clauses, where used, are intended to cover the structures described herein as performing the recited function and not only structural  
20 equivalents but also equivalent structures. Therefore, it is to be understood that the foregoing is illustrative of the present invention and is not to be construed as limited to the specific embodiments disclosed, and that modifications to the disclosed embodiments, as well as other embodiments, are intended to be included within the scope of the appended claims. The invention is defined by the following claims, with  
25 equivalents of the claims to be included therein.

## THAT WHICH IS CLAIMED IS:

1. A method for fabricating a plurality of sensors with electrodes on a flexible substrate so that the sensors each have at least one well associated therewith, comprising:
  - 5 providing a flexible substrate material layer having a surface area defined by a length and width thereof;
  - forming a plurality of sensors onto the flexible substrate material layer, each sensor comprising a metallic pattern defining at least one electrode;
  - disposing at least one photoimageable dry film coverlay sheet over the flexible
  - 10 substrate layer thereby sandwiching the sensors therebetween, the coverlay sheet having an associated thickness;
  - laminating the at least one coverlay sheet to the flexible substrate; and then, after laminating,
  - removing predetermined regions of the laminated coverlay sheet from the
  - 15 flexible substrate layer to define wells with a depth corresponding to the thickness of the at least one coverlay sheet.
2. A method according to Claim 1, wherein the well depth is at least about 100 $\mu$ m.
- 20 3. A method according to Claim 2, wherein the well depth is at least about 250 $\mu$ m.
4. A method according to Claim 1, wherein the well depth is in the range of about 1-20 mils.
- 25 5. A method according to Claim 1, wherein the at least one coverlay sheet comprises first and second coverlay sheets, said wherein, said method is carried out such that said first coverlay sheet is laminated to the flexible substrate material layer before said second coverlay sheet is disposed over the first laminated coverlay sheet,
- 30 and then said second coverlay sheet is positioned over said first laminated coverlay sheet and then laminated to said first coverlay sheet and said underlying flexible

substrate material layer, and wherein said removing step comprises removing corresponding overlying portions of both of the first and second coverlay sheets such that the well depth is defined by the combined thickness of the first and second coverlay sheets.

5

6. A method according to Claim 1, further comprising curing the at least one laminated coverlay sheet after said removing step.

7. A method according to Claim 1, wherein the sensors are configured so as to occupy a major portion of the surface area of the flexible substrate material layer such that the sensors are positioned on the flexible substrate material layer in a density of at least about 4 sensors per in<sup>2</sup> when measured over an area of at least about 121 in<sup>2</sup>.

8. A method according to Claim 1, wherein said removing step comprises:

selectively exposing predetermined regions of the at least one coverlay sheet excluding the region corresponding to the wells to ultraviolet light; and then introducing a liquid solution thereon to remove the portion of the at least one coverlay sheet overlying the exposed regions thereon to form the wells.

9. A method according to Claim 1, further comprising evacuating the substrate layer and coverlay sheet during laminating to remove air positioned between the coverlay sheet and the substrate layer.

25

10. A method according to Claim 8, wherein the metallic pattern for each sensor includes a pair of spaced apart metal bond pads, an interdigitated digital array (IDA), and a pair of metallic connecting traces, each of which extends between a respective bond pad and terminates into a respective opposing side of the IDA, and wherein the removing step exposes the metallic pattern associated with the bond pads and at least a portion of the IDA.

30

11. A method according to Claim 10, wherein said removing step removes portions of the at least one coverlay sheet overlying the metallic bond pads and a portion of the IDA but does not remove portions of the laminated coverlay sheet over the connecting traces about a major portion of the distance between the bond pads and the IDA for electric insulation.

12. A method according to Claim 1, wherein said providing, forming, disposing, laminating, and removing steps are carried out in a serially successive and substantially continuous manner.

10

13. A method according to Claim 1, further comprising curing the coverlay sheet after said removing step, and wherein said providing, forming, disposing, laminating, removing, and curing steps are repeated in an automated manner by processing the material as a continuous length through each of said steps, the continuous length being sufficient to roll onto a reel.

15

14. A method according to Claim 1, wherein said at least one coverlay sheet comprises at least first and second coverlay sheets, each having a different thickness.

20

15. A method according to Claim 14, wherein one of said first and second coverlay sheets is formed of a first photosensitive material, and the other is formed of a second different photosensitive material.

25

16. A method according to Claim 1, wherein the sensors are arranged on said substrate layer such that they define aligned columns and rows, the columns having a first quantity of sensors and the rows having a second quantity of sensors.

17. A method according to Claim 16, wherein said disposing step comprises conveying a continuous length of coverlay sheet material.

30

18. A method according to Claim 17, wherein said laminating step comprises:

conveying a continuous length of coverlay sheet material;  
conveying a continuous length of flexible substrate material with the metallic  
pattern formed thereon;

directing the coverlay sheet material and the flexible substrate material to meet  
5 such that the metallic pattern is sandwiched between the coverlay sheet and the  
flexible substrate material layer; and  
pressing the coverlay sheet and the flexible substrate material layer together.

19. A method according to Claim 1, wherein the at least one coverlay sheet  
10 includes two layers, a floor layer of a photoimageable dry film material and a ceiling  
layer disposed thereon.

20. A method according to Claim 1, wherein said removing step is carried  
out in a continuous or semi-continuous automated manner.

15

21. An array of flexible sensors, comprising:  
a flexible substrate layer having opposing primary surfaces;  
an electrode layer disposed as a repetition of metallic electrically conductive  
patterns on one of the primary surfaces of the first substrate layer, the metallic pattern  
20 corresponding to a desired electrode arrangement for a respective sensor; and  
a first coverlay sheet layer comprising a flexible photoimageable or  
photodefineable dry film material having an associated thickness overlying and  
laminated to said flexible substrate layer to sandwich said electrode layer  
therebetween, said first coverlay sheet layer having a plurality of photodefined apertures  
25 formed therein, said photodefined apertures defining a well for each of the sensors on  
the flexible substrate, wherein the wells have a depth corresponding to the thickness  
of the coverlay sheet layer.

22. An array according to Claim 21, further comprising a second coverlay  
30 sheet layer having a thickness overlying and secured to said first coverlay sheet layer,  
said second coverlay sheet layer having a plurality of photodefined apertures formed  
therein, the apertures corresponding to the apertures in the first coverlay sheet layer,  
wherein the wells have a depth corresponding to the combined thickness of the first

and second coverlay sheet layers, wherein the coverlay sheet is directly laminated to the underlying flexible substrate and is void of an added amount of adhesive positioned intermediate thereof.

5           23.     An array according to Claim 22, wherein the sensors are arranged in a pattern having a density of at least about 4 sensors per square inch when measured over about 122 square inches.

          24.     An array according to Claim 21, wherein the first and second coverlay  
10 sheet layer comprises a photosensitive dry film sheet, each having a different thickness.

          25.     An array according to Claim 21, wherein the array is arranged such  
that said sensors are sufficiently spaced apart to enable separation thereof into a  
15 plurality of discrete flexible sensors, and wherein, when separated and in operation, the sensor well is configured to receive a sample quantity of a predetermined biomaterial therein.

          26.     An array according to Claim 21, wherein the coverlay sheet layer has a  
20 thickness of at least about 1-10 mils.

          27.     An array according to Claim 21, wherein the coverlay sheet layer has a thickness of between about 5-20 mils.

25           28.     An array according to Claim 22, wherein said first and second coverlay sheets have substantially the same thickness.

          29.     An array according to Claim 22, wherein one of said first and second  
coverlay sheets is formed of a first photosensitive material, and the other is formed of  
30 a second different photosensitive material

          30.     A flexible sensor, comprising:  
a flexible substrate layer;

an electrode layer comprising a conductive pattern of metal disposed onto one of the primary surfaces of the flexible substrate layer; and

5 a first flexible photosensitive coverlay sheet layer overlying the second electrode layer and laminated to the electrode layer and the substrate layer so as to lack an intermediately positioned adhesive, wherein the first flexible coverlay sheet layer has a well formed therein, the well having a depth of at least about 1-10 mils.

31. A flexible sensor according to Claim 30, wherein said electrode layer conductive pattern includes two spaced apart bond pads, two connecting traces of  
10 which each terminates into an IDA located within the well.

32. A flexible sensor according to Claim 30, wherein, in operation, the well is adapted to receive a quantity of biofluid to generate an electrochemical response signal which is then transmitted to the bond pads.  
15

33. A flexible sensor according to Claim 30, wherein the first coverlay sheet material well is photodefined.

34. A flexible sensor according to Claim 30, wherein the first coverlay  
20 sheet layer has a thickness of between about 5-20 mils.

35. A flexible sensor according to Claim 30, further comprising a second coverlay sheet layer having a thickness overlying and secured to said first coverlay sheet layer, said second coverlay sheet layer having a plurality of apertures formed  
25 therein, the apertures corresponding to the apertures in the first coverlay sheet layer, wherein the wells have a depth corresponding to the combined thickness of the first and second coverlay sheet layers.

36. A flexible sensor according to Claim 35, wherein said first and second  
30 coverlay sheets have a different thickness.



37. A flexible sensor according to Claim 35, wherein one of said first and second coverlay sheets is formed of a first photosensitive material, and the other is formed of a second different photosensitive material.

5           38. A flexible sensor according to Claim 30, wherein the first dry film coverlay sheet is directly laminated to the underlying electrode and/or substrate layer so as to be void of adhesive therebetween.

10           39. A system for concurrently fabricating a plurality of flexible sensors, with electrodes on a flexible substrate so that the sensors each have at least one well associated therewith, comprising:

          means for providing a flexible substrate material layer having a surface area defined by a length and width thereof;

15           means for forming a plurality of sensors onto the flexible substrate material layer, each sensor comprising a metallic pattern defining at least one electrode;

          means for disposing at least one photoimageable dry film coverlay sheet over the flexible substrate layer thereby sandwiching the sensors therebetween, the coverlay sheet having an associated thickness;

20           means for laminating the at least one coverlay sheet to the flexible substrate; and then, after laminating, and

          means for removing predetermined regions of the laminated coverlay sheet from the flexible substrate layer to define wells with a depth corresponding to the thickness of the at least one coverlay sheet.

25           40. A system according to Claim 39, wherein said means for laminating is carried out with:

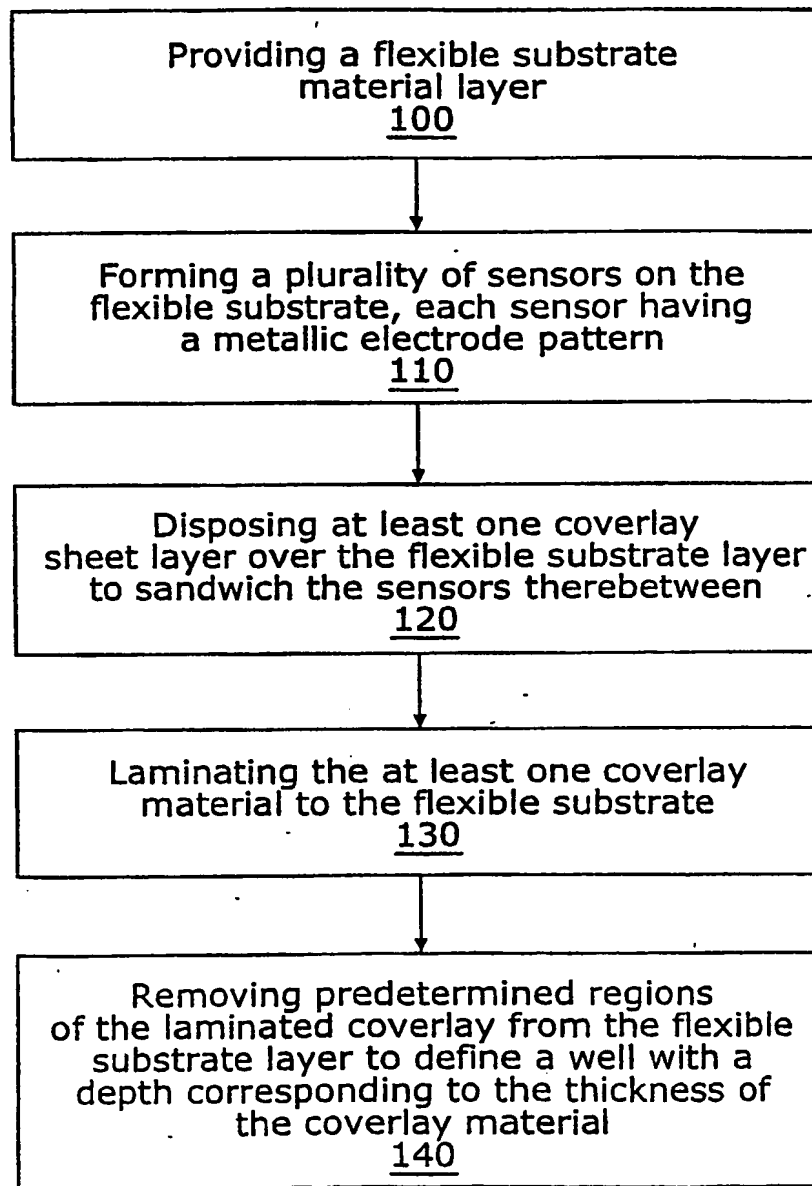
          means for conveying a continuous length of coverlay sheet material;

          means for conveying a continuous length of flexible substrate material with the metallic pattern formed thereon;

30           means for directing the coverlay sheet material and the flexible substrate material to meet such that the metallic pattern is sandwiched between the coverlay sheet and the flexible substrate material layer; and

means for pressing the coverlay sheet and the flexible substrate material layer together.

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FIG. 1

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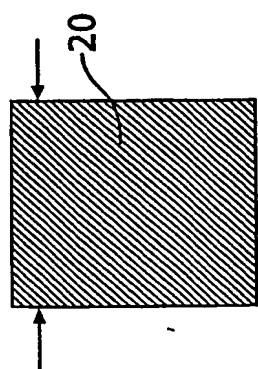
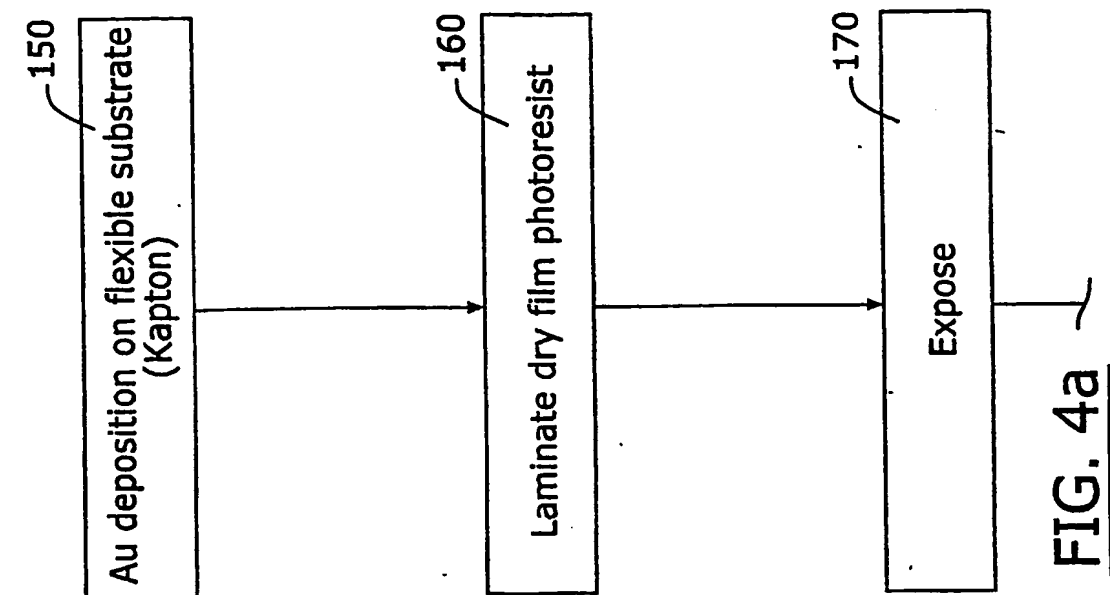


FIG. 3a

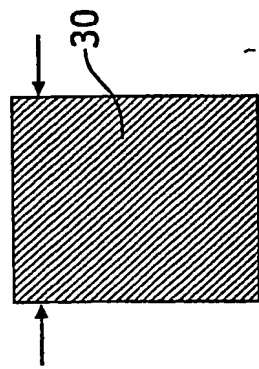


FIG. 3b

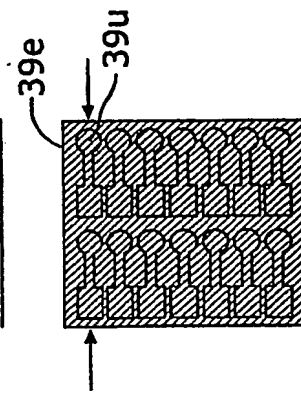


FIG. 3c

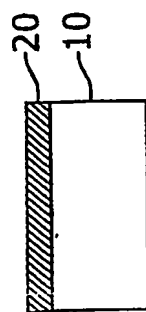


FIG. 2a

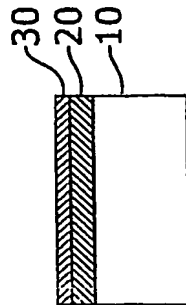


FIG. 2b

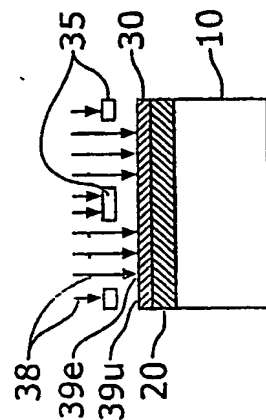
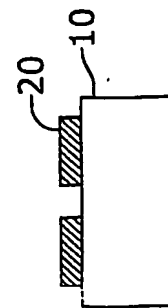
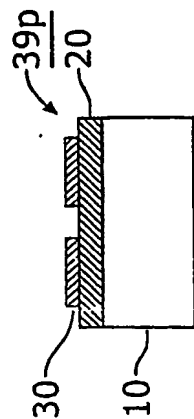
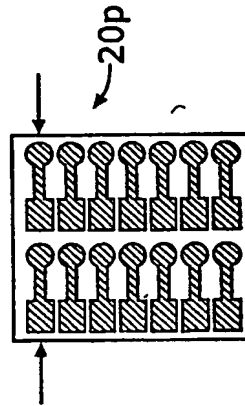
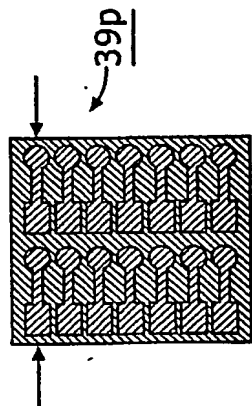
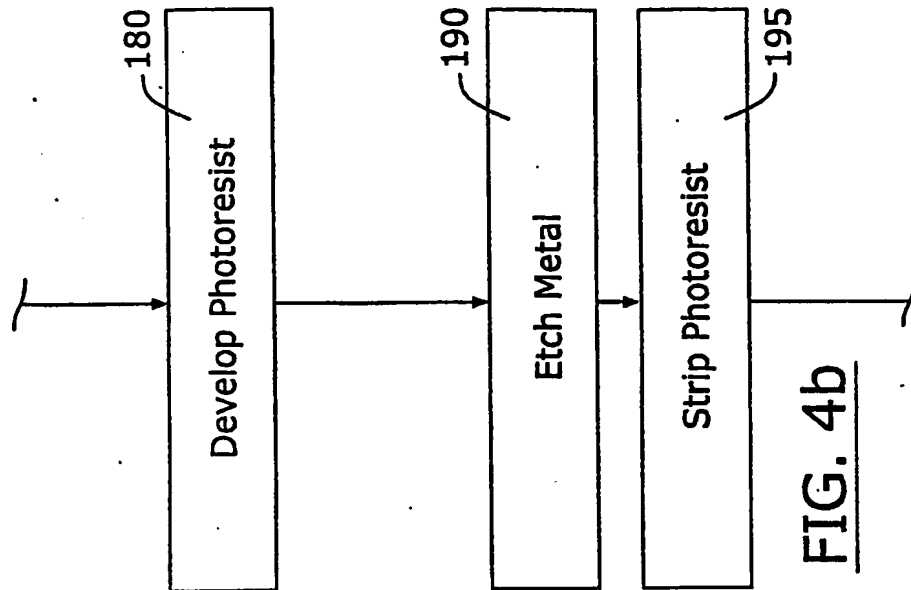


FIG. 2c

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4/10

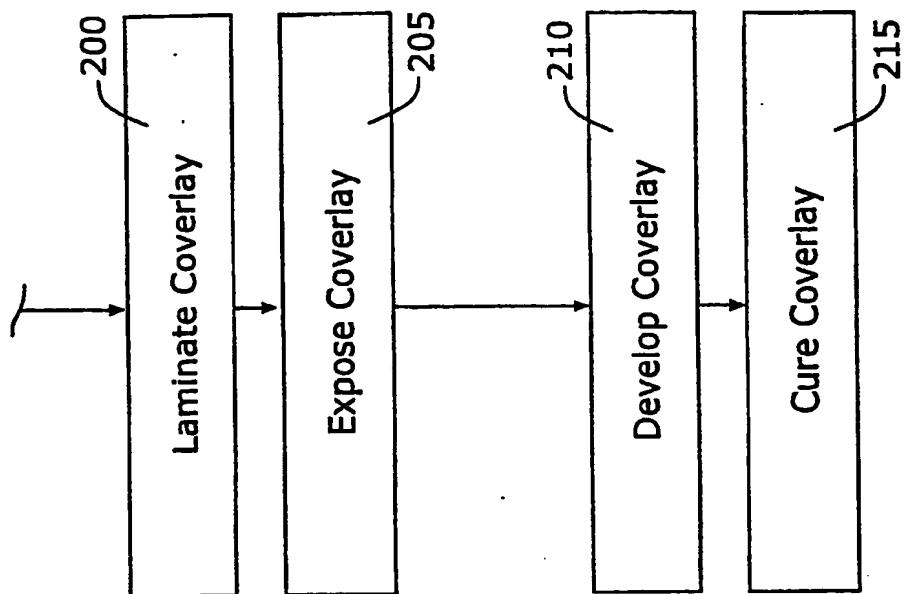


FIG. 4c

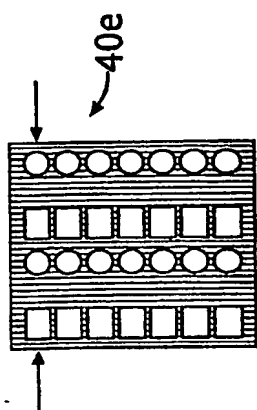


FIG. 3f

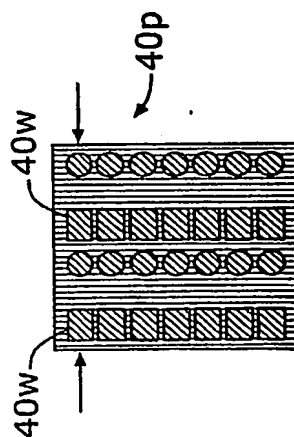


FIG. 3g

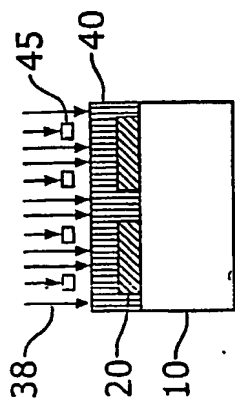


FIG. 2f

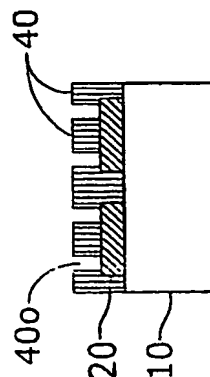


FIG. 2g

5/10

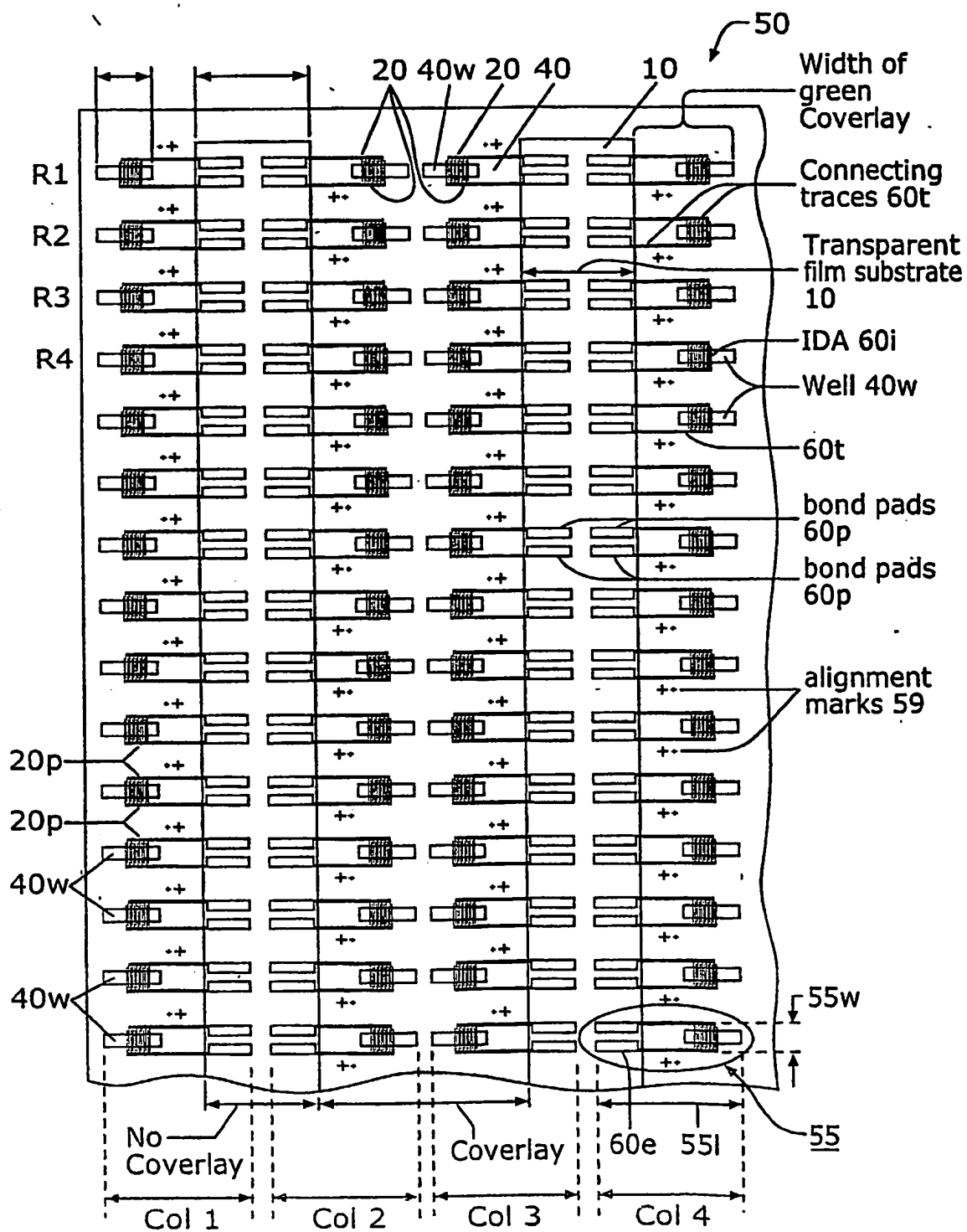
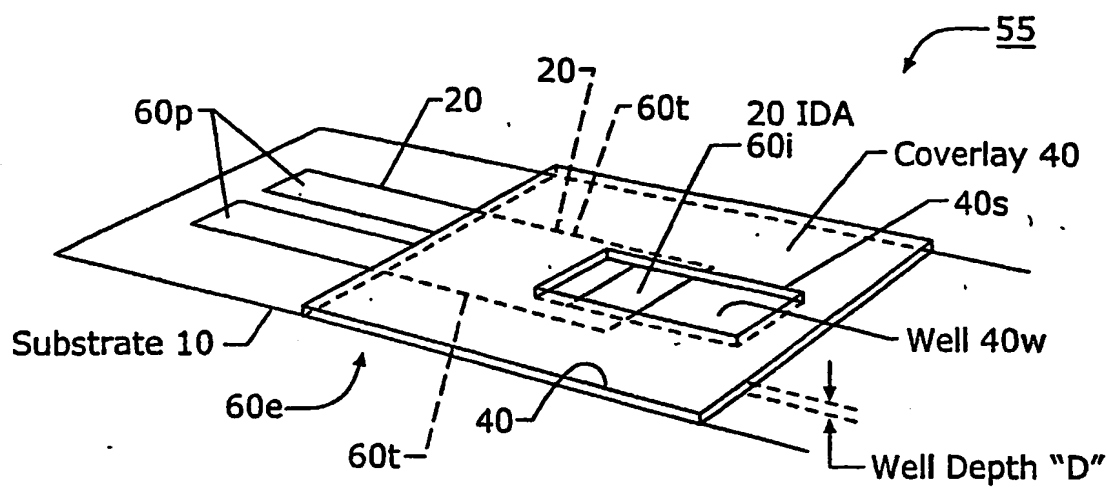


FIG. 5

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**FIG. 6a**

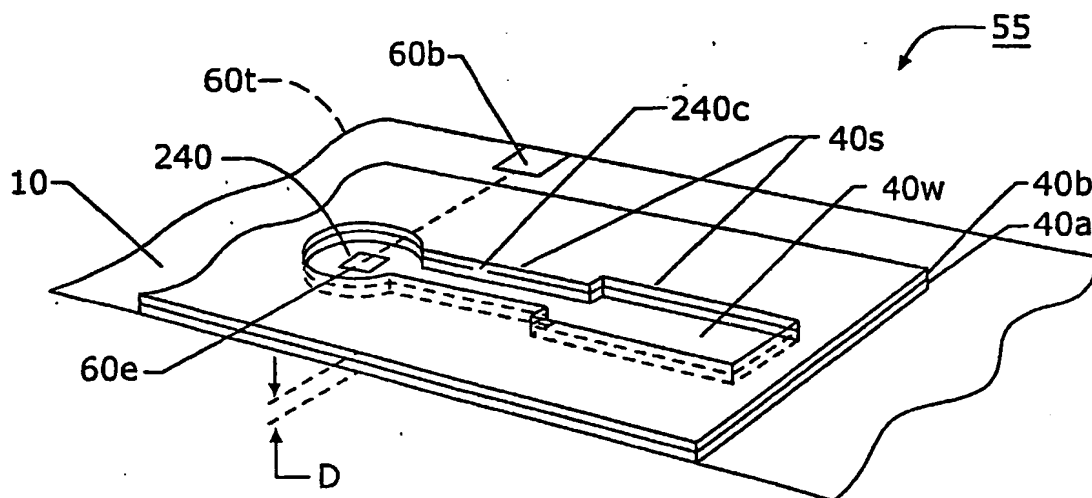


FIG. 6b



7/10

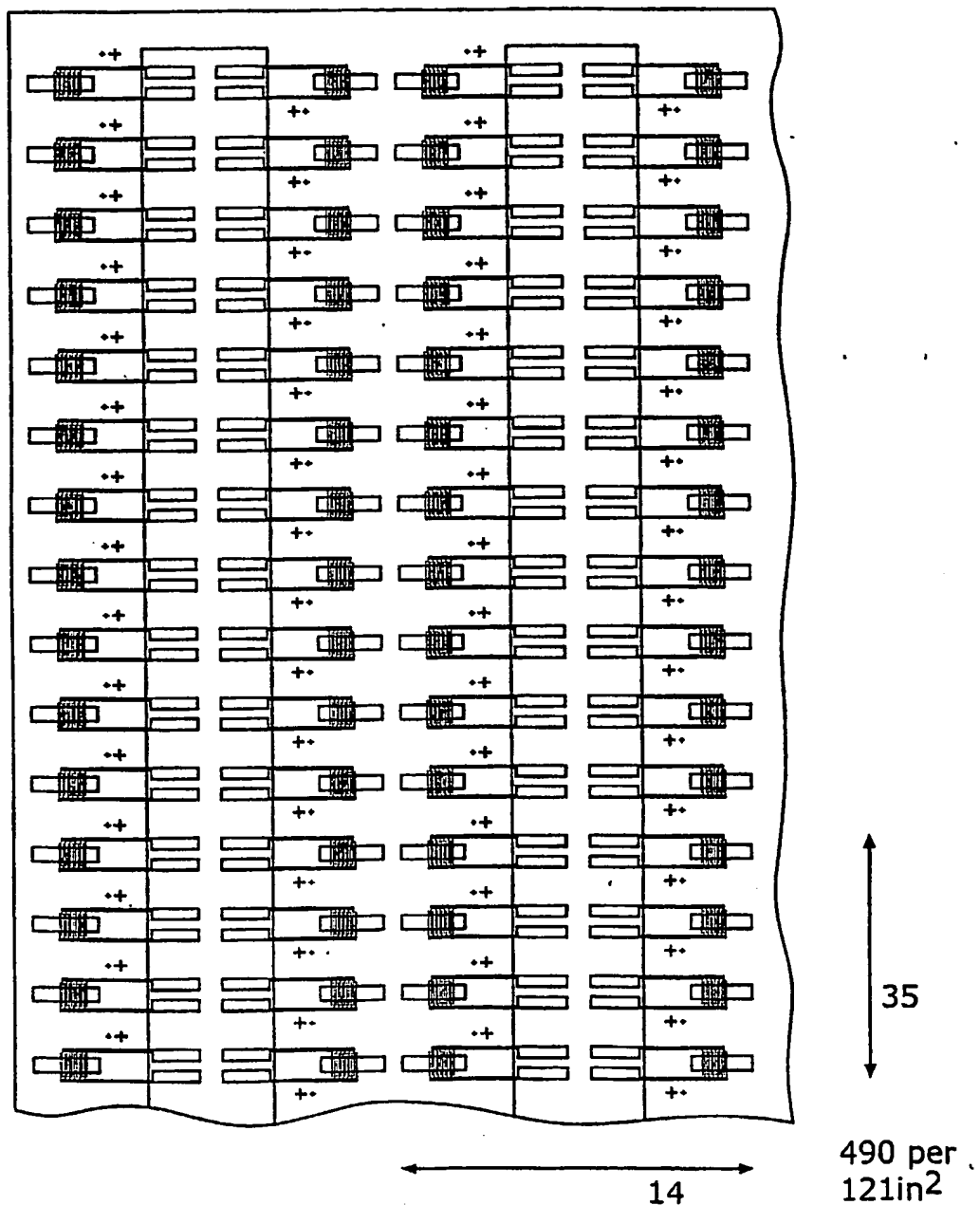


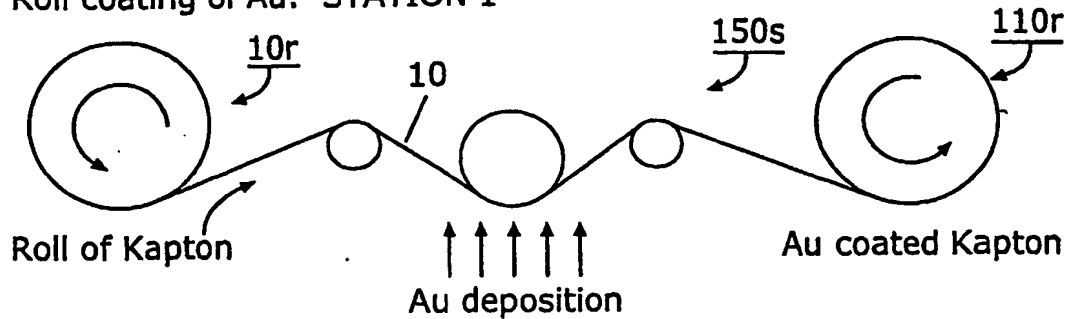
FIG. 7

SUBSTITUTE SHEET (RULE 26)

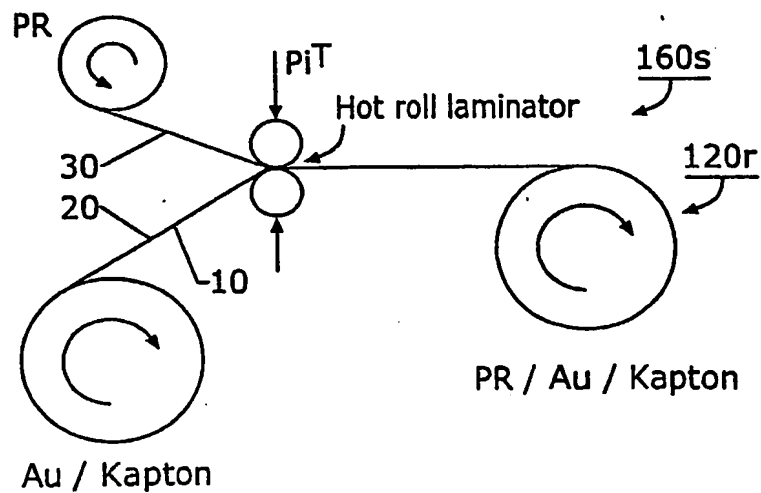
8/10

## Continuous process using Au coated Kapton foil

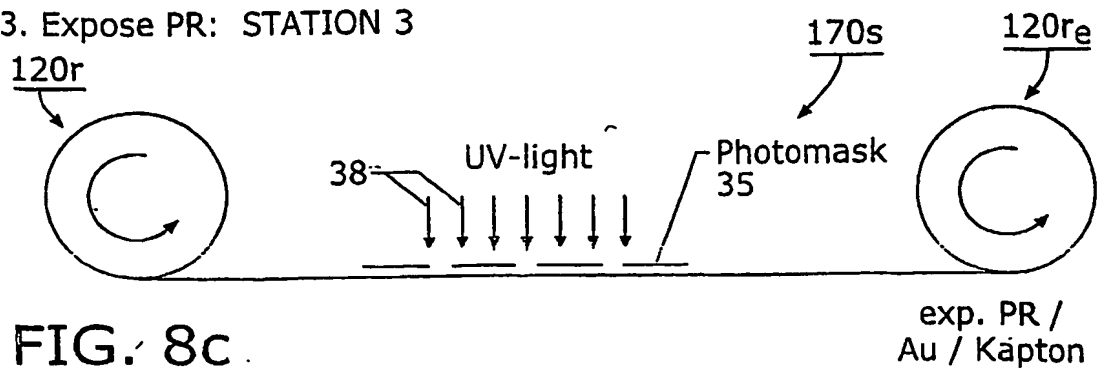
## 1. Roll coating of Au: STATION 1

FIG. 8a

## 2. Laminate dry photoresist: STATION 2

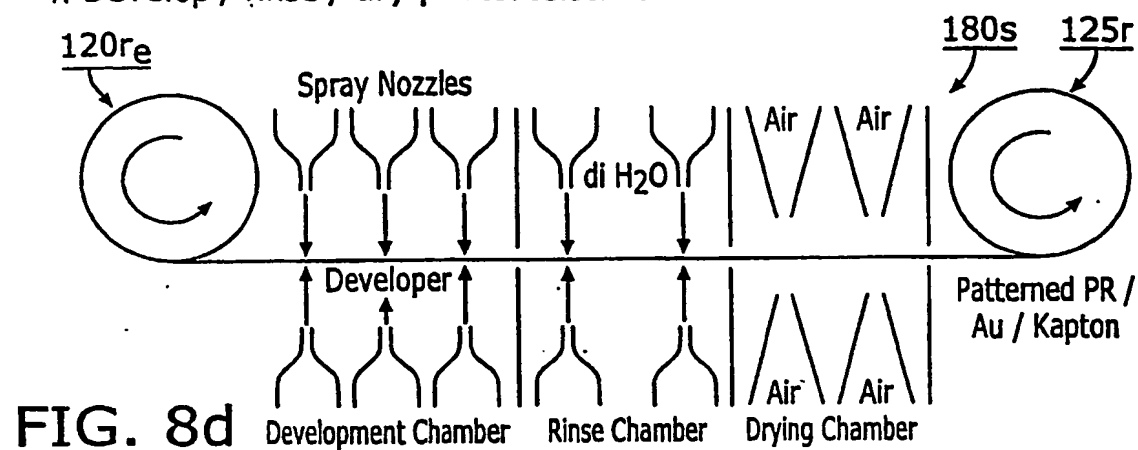
FIG. 8b

## 3. Expose PR: STATION 3

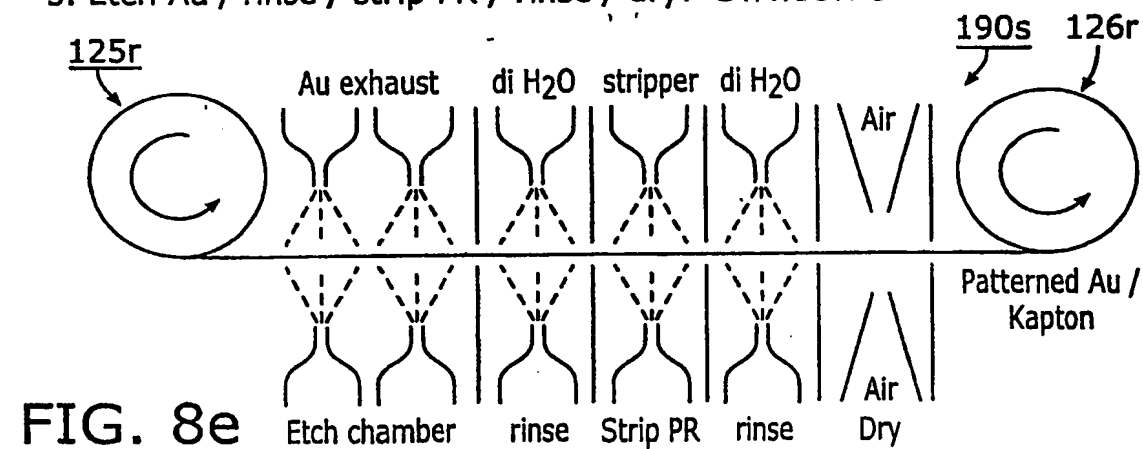
FIG. 8c

9/10

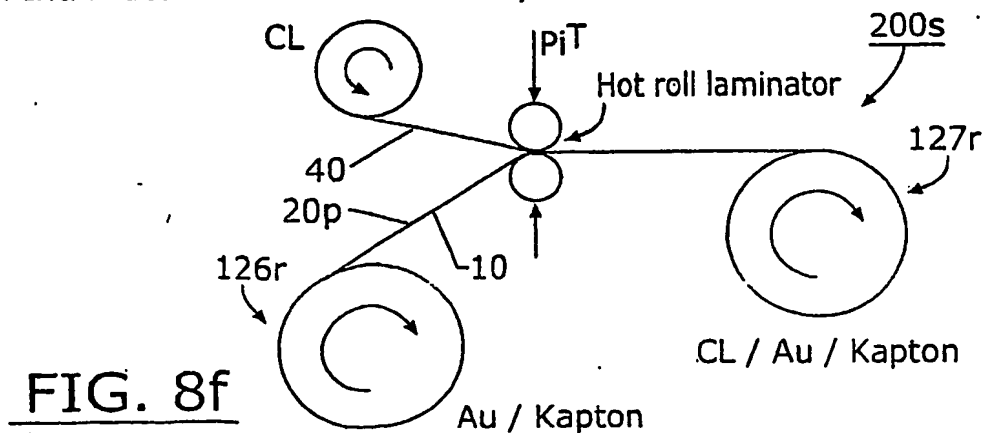
## 4. Develop / rinse / dry photoresist: STATION 4



## 5. Etch Au / rinse / strip PR / rinse / dry: STATION 5

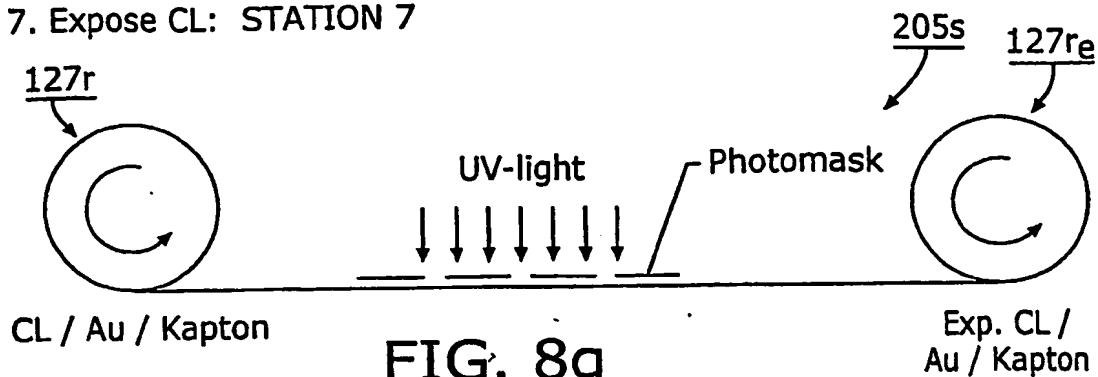


## 6. Introduce and laminate coverlay: STATION 6

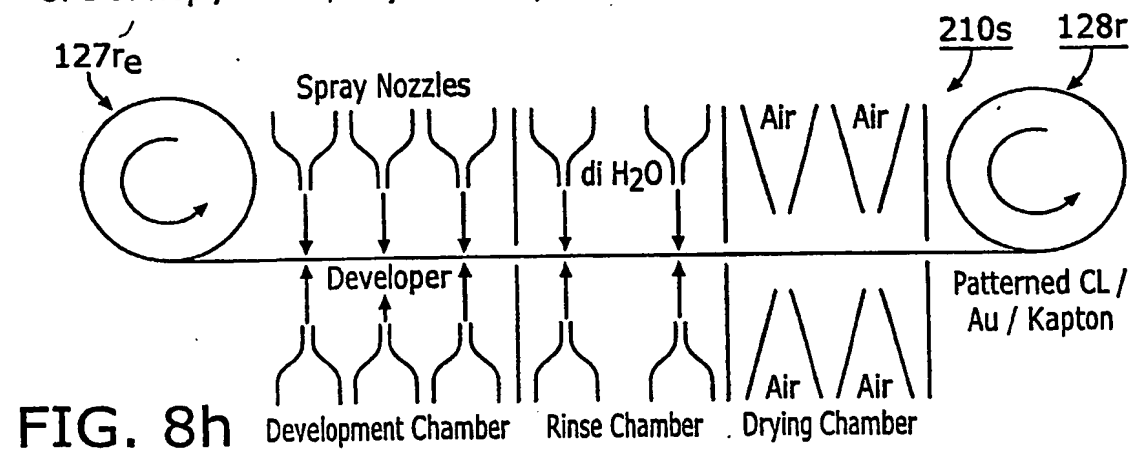


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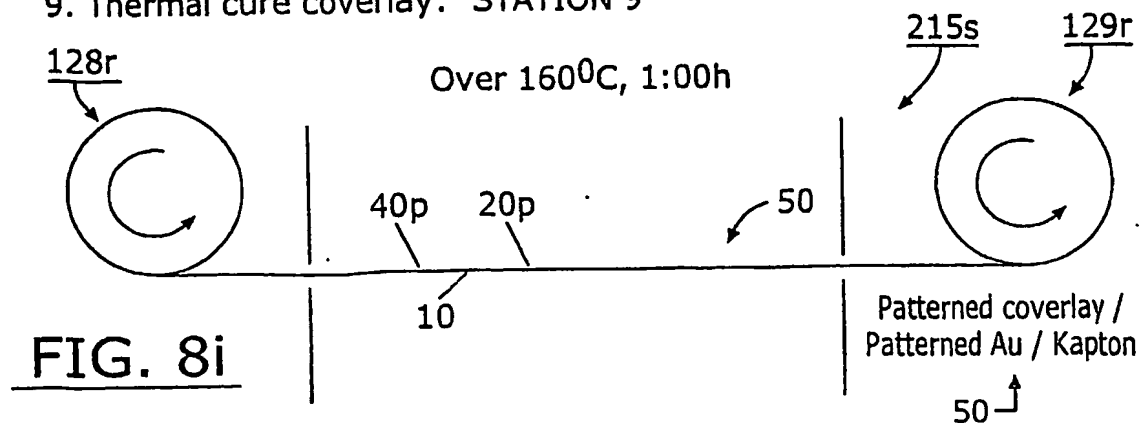
## 7. Expose CL: STATION 7

**FIG. 8g**

## 8. Develop / rinse / dry coverlay: STATION 8

**FIG. 8h**

## 9. Thermal cure coverlay: STATION 9

**FIG. 8i**

# INTERNATIONAL SEARCH REPORT

International application No.

PCT/US02/36774

## A. CLASSIFICATION OF SUBJECT MATTER

IPC(7) : C03C 15/00, 25/68; G01N 27/26; C12M 1/34, 03/00; B44C 1/165

US CL : 216/33, 204/422, 435/287.8, 156/230

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 216/33, 36; 204/422, 403.03, 403.13; 435/287.1-287.03, 287.08, 287.09; 156/230, 238, 73.1

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)  
Please See Continuation Sheet

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	US 6,103,033 A (SAY et al.) 15 August 2000 (15.8.2000), column 24, lines 46-57.	1-20
Y	US 5,437,999 A (DIEBOLD et al.) 01 June 1995 (01.6.1995), column 9, lines 4-63.	30-38

☐ Further documents are listed in the continuation of Box C.

☐ See patent family annex.

\* Special categories of cited documents:

"A" document defining the general state of the art which is not considered to be of particular relevance

"E" earlier application or patent published on or after the international filing date

"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art

"&" document member of the same patent family

Date of the actual completion of the international search

10 March 2003 (10.03.2003)

Date of mailing of the international search report

15 APR 2003

Name and mailing address of the ISA/US

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**INTERNATIONAL SEARCH REPORT**

PCT/US02/36774

**Continuation of Item 4 of the first sheet:**

The title is not descriptive and too long.

The new title is :Biomedical Electrochemical Sensor Array and Method of Fabrication

**Continuation of B. FIELDS SEARCHED Item 3:**

**EAST**

search terms: sensor, flexible, laminate, electrochemical, biomedical, remove, etch, continuous, photosensitive, resist, pattern, electrode, traces, web, roll